JUL 2 5 700 B

INFORMATION DISCLOSURE STATEMENT BY APPLICANT

(use as many sheets as necessary)

Application Number	09/859,604	
Filing Date	May 17, 2001	
First Named Inventor	Jack R. Wands	
Group Art Unit	Not yet assigned	
Examiner Name	Not yet assigned	
Attorney Docket Number	21486-032CIP	

Exam Initials	Cite No.	U.S. Patent Document No.	Issue Date	Name of Patentee(s) or Applicant(s)	Class	Sub Class	Filing Date If Appropriate	
KAC	AA	6,165,464	12/26/00	Hudziak, et al.	424	142.1		
		2 (2) (2) (2) (2) (2) (2) (2) (2) (2) (2		FOREIGN PATENT DOCUMENTS	Carry			
Exam Initials	Cite No.	Foreign Patent Do Office Number	ocument	Name of Patentee(s) or Applicant(s)	Date of Pu	blication	Translation Yes No	

Exam Initials	Cite No.	Name of Author, Title (when appropriate), Publication, Volume, Page(s), Date, Etc.							
KAC	CA	Aster J. et al., "Functional Analysis of the TAN –1 Gene, a Human Homolog of Drosphilia Notch", Cold Spring Harb. Symp. Uant. Biol., 1994; 59:125-36.							
	СВ	Baron et al., "Insulin Receptor-Induced Phosphorylation of Cellular and Synthetic Substrates Is Regualted by the Receptor β-Subunit C-Terminus", Endocrinology, 1996 August; 137(8) 3416-23.							
	СС	Boder, et al., "Directed Evolution of Antibody Fragments with Monovalent Femtomolar Anitgen-Binding Affinity", PNAS, September 26, 2000, vol. 97 (20), 10701-10705.							
	CD	Branch, "A hitchhikers guide to antisense and nonantisense biochemical pathways", Hepatology, Vol. 24, 1996, pp. 1517-1529.							
	CE	Broaddus et al., "Strategies for the design and delivery of antisense oligonucleotides in central nervous system", Methods in Enzymology, 2000, Vol. 314, pp. 121-135.							
	CF	Capobianco et al., "Neoplastic Transformation by Truncated Alleles of Human NOTCH1/TAN1 and NOTCH2", 1997, Mol. Cell Biol. 17:6265-6273							
	CG	Czubayko et al., "Ribozyme-targeting Elucidates a Direct Role of Pleiotrophin in Tumor Growth", 1994, J. Biol. Chem. 269:21358-21363							
	СН	de la Monte et al., "Microtiter Immunocytochemical ELISA Assay," 1999, Biotechniques							
	CI	de la Monte SM, et al., "Differential Effects of Ethanol on Insulin-Signaling Through the Insulin Receptor Substrate-1", Alcohol Clin. Exp. Res., 1999 May; 23(5):770-7.							
	Cı	Dinchuk, et al., "Aspartyl β-Hydroxylase (Asph) and an Evolutionarily Conserved Isoform of Asph Missing the Catalytic Domain Share Exons with Junctin," (2000) 275, 39543-54							
	CK	Foote, et al., "Breaking the Affintiy Ceiling for Antibodies and T Cell Receptors", PNAS, September 26, 2000, vol.97(20), 10679-10681.							
	CL	GENBANK Accession No. S83325							
	СМ	GENBANK Accession No. JS0670							
/	CN	GENBANK Accession No. NM 005544							
V	со	Ghose et al., "Immunochemical Techniques, part F, Conventional Antibodies Fc Receptors and Cytotoxicity", Methods in Enzymology, Vol. 93, 326-327, 1983							
Examine Signatur	-	Jane J. Ganelle Date Considered 6/13/03							

EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered.

Include copy of this form with next communication to applicant.



Modified	Form 1449	9/PTO		Application Number	09/859	,604			
				Filing Date	May 1	May 17, 2001			
	INI	FORMATION D	ISCLOSURE	First Named Inventor	First Named Inventor Jack R. Wands				
	STATEMENT BY APPLICANT			Group Art Unit	Not ye	Not yet assigned			
				Examiner Name	Not ye	Not yet assigned			
		(use as many sheets a	as necessary)	Attorney Docket Number	21486-032CIP				
er av Hajari s		oledio anis iri		U.S. PATENT DOCUMENTS					
Exam Initials	Exam Cite U.S. Patent Issue Date Name of Pa			Name of Patentee(s) or Applicant(s)	Class	Sub Class	Filing Date If Appropriate		

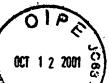
Exam Initials	Cite No.	Name of Author, Title (when appropriate), Publication, Volume, Page(s), Date, Etc.							
(AL	СР	Gronke et al., "Aspartyl β-hydroxlase: In vitro hydroxylation of a synthetic peptide based on the structure of the first growth factor-like domain of human factor IX", Proc. Natl. Acad. Sci. USA, Vol. 86, May 1989, pp. 3609-3613.							
	CQ	Gronke et al., "Partial Purification and Characterization of Bovine Liver Aspartyl β-Hydroxylase", 1990, J. Biol. Chem. 265:8558-8565							
	CR	Hagemeijer, "Cytogenetics and oncogenes", Leukemia, November 1992, Vol. 6, Suppl. 4, pp.16-18.							
	CS	Hansen T. et al., "Inhibition of insulin receptor phosphorylation by peptides derived from major histocompatability complex class I antigens,", Proc. Natl. Acad. Sci. USA 1989 May ;86 (9):3123-6.							
	CT 4	Higgins et al., "Fast and sensitive multiple sequence alignments on a microcomputer", 1989, CABIOS 5(2):151-153							
	CU	Jia et al., "A fully active atalytic domain of bovine aspartyl (asparaginyl) β-hydroxylase expressed in Escherichia Coli", Proc. Natl. Acad. Sci. USA 1994 July 19;91(15):7227-31.							
	cv	Jia et al., "cDNA Cloning and Expression Bovine Aspartyl (Asparginyl) β-Hydroxylase", 1992, J. Biol. Chem. 267:14322-14327							
	CW	Jones et al., "Antibodies for targeting gene therapy", Advanced Drug Delivery Reviews, Apr. 1998, Vol. 31, pp. 153-170.							
	CX	Kelley et al., "Mutations Altering the Structure of Epidermal Growth Factor-like Coding Sequences at the Drosophilia Notch Locus, Cell., Nov. 20, 1997, Vol. 51, p. 539-548.							
	CY	Kieke, et al., "Selection of Functional T Cell Receptor Mutants From a Yeast Surface-Display Library", Proc. Natl, Acad. Sci, USA, May 1999, Vol. 97, 5651-5656							
	CZ	Kobayashi et al., "Reversal of Drug Sensitivity in Multidrug-Resistant Tumor Cells by an MDR1 (PGY1) Ribozyme", 1994, Cancer Res. 54:1271-1275							
	CAA	Korioth et al., "Cloning and characterization of the human gene encoding aspartyl β-hydroxylase", Gene, 150, 1994, 395-399							
	CAB	Lam K., et al., "The Phosphatidylinositol 3-Kinase Serine Kinase Phosporylyates IRS-1", J. Biol. Chem., 1994 August 12:269(32):20648-52							
	CAC	Lardelli et al., "The Novel Notch homologue mouse Notch 3 lacks specific epidermal growth factor-repeats and is expressed in proliferating neurothelium", Mechanisms of Development, 46 (1994) 123-136.							
	CAD	Lavaissiere et al., "Overexpression of Human Aspartyl (asparaginyl) β-Hydroxylase in Heptocellular Carcinoma and Cholangiocarcenoma", 1996, J. Clin. Invest. 98:1313-1323							
$\sqrt{}$	CAE	Lecka-Czernik et al., "An Overexpressed Gene Transcript in Senescent and Quiescent Human Fibrolasts Encoding a Novel Protein in the Epidermal Growth Factor-Like Repeat Family Stimulates DNA Synthesis", Molecular and Cellular Biology, Vol. 15, Jan. 1995, p. 120-128.							
Examine Signatur	e 4	Faren J. Canella Date Considered 6/13/03							

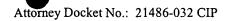
EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

Francis City II C Detroit				ame of Patentee(s) or Applicant(s)	Class	Sub Class	Filing Date If Appropriate		
		(use as many sheets	• •	Attorney Docket Number U.S. PATENT DOCUMENTS	21486-032CIP				
				Examiner Name	Not y	Not yet assigned			
	STATEMENT BY APPLICANT			Group Art Unit	Not y	Not yet assigned			
	INI	FORMATION D	ISCLOSURE	First Named Inventor	Jack l	R. Wands			
				Filing Date	May	17, 2001			
Modified I	Form 1449	P/PTO		Application Number	09/85	9,604			

Exam Initials	Cite No.	Name of Author, Title (when appropriate), Publication, Volume, Page(s), Date, Etc.							
KAL	CAF	Levy-Toledano R. et al., "Investgation of the mechanism of the dominant negative effect of mutations in the tyrosine kinase domain of the insulin receptor, EMBO J. 1994 Feb 15;13(4):825-42.							
	CAG	Li et al., "Modulation of Insulin Receptor Substrate-1 Tyrosine Phosphoryllation by an Akt/Phospatidylinositol 3-Kinase Pathway", J. Biol. Chem., 1999 Apr 2;274 (14):9351-6.							
	CAH	Mahieu et al., "Construction of a Ribozyme Directed Against Human Interleukin-6 mRNA; Evaluation of Its Catalytic Activity In Vitro and In Vivo", 1994, Blood 84:3758-65							
	CAI	Marasco et al., "Design, intracellular expression, and activity of a human anti-human immunodeficiency virus type-1 gp120 single-chain antibody" 1993, Proc. Natl., Acad. Sci. USA 90:7889-7893. (1998) 454-456							
	CAJ	Marasco et al., "Intrabodies: turning the humoral immune system outside in for intracellular immunization", 1997, Gene Therapy 4:11-15							
	CAK	McGinnis et al., "The five cysteine residues located in the active site region of bovine aspartyl β-hydroxylase are not essential for catalysis", Biochemica et Biophysica Acta 1387							
	CAL	McGinnis et al., "Site-Directed Mutagenesis of Residues in a ConservedRegion of Bovine Aspartayl β-Hydroxylase: Evidenc That Histidine 675 Has a Role in Binding Fe", Biochemistry 1996, 35, 3957-3962.							
	CAM	Rozan F., Pollack, M., "Inhibition of insulin-like growth factor I receptor signaling by the vitamin D analogue EB 1089 in MCF-7 breast cancer cells: A role for insulin-like growth factor binding proteins", Int. J. Oncol., 1999, Sept. 15(3):589-94.							
	CAN	Song et al., "Proteolytic release and nuclear translocation of notch-1 are induced by presenilin-1 and impaired by pathogenic presenilin-1 mutations", 1999, Proc. Natl. Acad. Sci U.S.A. 96:6959-6963							
	CAO	Sullivan et al., "Development of Ribozymes for Gene Therapy", 1994, J. Invest. Derm. 103:85S-89S							
	CAP	Treves, et al. "Molecular Cloning, Expression, Functional Characterization, Chromosomal Localization, and Gene Structure o Junctate, a Novel Integral Calcium Binding Protein of Sarco(endo)plasmic Reticulum Membrane," (2000) 275, 39555-68							
	CAQ	Van de Poll ML et al., "Identification of the Minimal Requirements for Binding to the Human Epidermal Growth Factor (EGF) Receptor Using Chimeras of Human EGF and an EGF Repeat of Drosphilia Notch", J. Biol. Chem. 1998 June 26;273(26):16075-81.							
1 /	CAR	Wang et al., "Bovine Liver Aspartyl β-Hydroxylase", 1991, J. Biol. Chem. 266:14004-14010							
	CAS	Zhang et al., "Development of a Carbon Dioxide-Capture Assay in Microtiter Plate for Aspartyl β-Hydroxylase", 1999, Anal. Biochem. 271:137-142							
V									
Examiner Signature		Harm J. Ganelle Date Considered 6/13/03							

EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.





LIST O	FORM PTO-1449(Modified) LIST OF PATENTS AND PUBLICATIONS FOR APPLICANT'S INFORMATION DISCLOSURE STATEMENT				ATTY. DOCKET NO. SERIAL NO. 21486-032 CIP 09/859,604				
	"Diagnosis and Treatment of Malignant Neoplasms"			FIRST NAMED APPLICANT: Wands			9 2001 9 2001		
					FILING DATE: May 17, 2001		GR0 1642	OUP:	9 2001
	·			U.S. PATE	NT DOCUMENTS	-			- 53 -
Exam Init	Ref Des	Document No.	Date	Name		Cla	ss	Sub Class	FILIT DATE If Appropriate
KAC	A2	US 6,166,176	12/26/00	Radosevich			1		03/17/98
			F	OREIGN PA	TENT DOCUMENTS				
		Country & Doc. No. (11)	Pub. Date (43)			Cla	ss	Sub Class	Translation Yes No
									
KAC	C46 C47 C48 C49 C50 C51 C52 C53	De La Monte Ghose, T. et a Gual, P. et al Ince, N. et al Ince, N. et al Nishimaki, H	et al. <u>Curren</u> e, S. et al. (1 al. (1983). . (1996). En . (1997). Ho . (2000). Co I. et al. (199 I. et al. (199	t Protocols 999). Mod Methods in ndocrinology epatology? ancer Rese 9). Jikeika 7). Gastro	e, Pertinent Pages, Publication, E in Molecular Biology, V dern Pathology 12: 170A Enzymology 93: 280-33 gy 137: 3416-2423. 26(4): 362A. arch 60: 1261-1266. di Medical Journal 46: 1 enterology 112(4)Supp.: June 5, 2001.	7ol. 2 19 3. 29-136	•	John W	iley & Sons.
·	U34	1 micriational	Scaron Rep	ori, 133ucu	Julio 2, 2001.				
applica U.S.C.	tion, S 120 (c	erial No,	filedtinuation-in-p	, since aban part, and div	iously cited by or submitted doned, and relied upon for isional applications). DATE CONSIDERED	an earlie	office er filin	in a prio	nder 35

EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw a line through citation if not in conformance and not considered.

Please include copy of this form with next communication to applicant